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| U.S. DEPARTMENT OF COMMERCE, PATENT AND TRADEMARK OFFICE   |   | DATE: March 2, 2006                                 |
| TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE<br>(DO/EO/US) CONCERNING A FILING UNDER 35 U.S.C. 371  |   | U.S. Application No. (if known)<br><b>10/570252</b> |
| INTERNATIONAL APPLICATION NO.:<br><b>PCT/JP2005/001055</b>   | INTERNATIONAL FILING DATE:<br><b>January 27, 2005</b> | PRIORITY DATE CLAIMED:<br><b>March 3, 2004</b>      |
| TITLE OF INVENTION: <b>THERAPEUTIC OR PREVENTIVE DRUG FOR OSTEOPOROSIS COMPRISING ISOTAXIRESINOL DERIVED FROM TAXUS YUNNANENSIS</b>  |   |   |
| APPLICANT(S) FOR DO/EO/US: <b>Shigetoshi KADOTA et al.</b>   |   |   |
| Applicant hereby submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:  |   |   |
| <ol style="list-style-type: none"> <li>1. <input checked="" type="checkbox"/> This is a FIRST submission of items concerning a filing under 35 U.S.C. 371.</li> <li>2. <input type="checkbox"/> This is a SECOND or SUBSEQUENT submission of items concerning a filing under 35 U.S.C. 371.</li> <li>3. <input checked="" type="checkbox"/> This is an express request to begin national examination procedures (35 USC 371(f)). The submission must include items (5), (6), (9) and (21) indicated below.</li> <li>4. <input checked="" type="checkbox"/> The US has been elected (Article 31).</li> <li>5. <input checked="" type="checkbox"/> A copy of the International Application as filed (35 U.S.C. 371(c)(2)):               <ol style="list-style-type: none"> <li>a. <input type="checkbox"/> is transmitted herewith (required only if not transmitted by the International Bureau).</li> <li>b. <input checked="" type="checkbox"/> has been transmitted by the International Bureau.</li> <li>c. <input type="checkbox"/> is not required, as the application was filed in the United States Receiving Office (RO/US)</li> </ol> </li> <li>6. <input checked="" type="checkbox"/> An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)).               <ol style="list-style-type: none"> <li>a. <input checked="" type="checkbox"/> is attached hereto.</li> <li>b. <input type="checkbox"/> has been previously submitted under 35 U.S.C. 154(d)(4).</li> </ol> </li> <li>7. <input checked="" type="checkbox"/> Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))               <ol style="list-style-type: none"> <li>a. <input type="checkbox"/> are transmitted herewith (required only if not transmitted by the International Bureau).</li> <li>b. <input type="checkbox"/> have been transmitted by the International Bureau.</li> <li>c. <input type="checkbox"/> have not been made; however, the time limit for making such amendments has NOT expired.</li> <li>d. <input checked="" type="checkbox"/> have not been made and will not be made.</li> </ol> </li> <li>8. <input type="checkbox"/> An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)).</li> <li>9. <input checked="" type="checkbox"/> An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).</li> <li>10. <input type="checkbox"/> An English language translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).</li> </ol> |   |   |
| ITEMS 11. TO 20. BELOW CONCERN OTHER DOCUMENT(S) OR INFORMATION INCLUDED:  |   |   |
| <ol style="list-style-type: none"> <li>11. <input checked="" type="checkbox"/> An Information Disclosure Statement under 37 CFR 1.97 and 1.98 together with the international search report, PTO/SB/08 and 4 docs.</li> <li>12. <input checked="" type="checkbox"/> An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.<br/>               ASSIGNEE NAME AND ADDRESS: <b>KOTOSUGI INC., Sagamihara-shi, Japan</b><br/>               Please publish the assignee data with the application.</li> <li>13. <input type="checkbox"/> A preliminary amendment.</li> <li>14. <input checked="" type="checkbox"/> An Application Data Sheet under 37 C.F.R. 1.76</li> <li>15. <input type="checkbox"/> A substitute specification.</li> <li>16. <input type="checkbox"/> A change of power of attorney and/or address letter.</li> <li>17. <input type="checkbox"/> A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 C.F.R. 1.821 - 1.825.</li> <li>18. <input type="checkbox"/> A second copy of the published international application under 35 U.S.C. 154(d)(4).</li> <li>19. <input type="checkbox"/> A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).</li> <li>20. <input type="checkbox"/> Other items or information:</li> </ol>   |   |   |

2006 MAR 21 P 4: 57  
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1AP20 Rec'd PCT/PTO 02 MAR 2006

ATTORNEY'S DOCKET NO.: 062043

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| U.S. APPLICATION NO. (if known)<br><b>10/1570252</b>   |              | INTERNATIONAL APPLICATION NO.<br>PCT/JP2005/001055 |  | DATE: March 2, 2006  |              |
| The following fees are submitted:  |              |  |  | CALCULATIONS   | PTO USE ONLY |
| 21. <input checked="" type="checkbox"/> Basic National Stage Fee \$300<br>22. <input checked="" type="checkbox"/> National Stage Examination Fee<br>If International preliminary examination report prepared by USPTO and all claims satisfy provisions of PCT Article 33(1)-(4) \$100<br>All other situations \$200<br><br>23. <input checked="" type="checkbox"/> National Stage Search Fee<br>Search fee (37 CFR 1.445(a)(2) has been paid on the international application to the USPTO as an International Searching Authority \$100<br>International Search Report prepared and provided to the Office \$400<br>All other situations \$500<br><br>ENTER APPROPRIATE BASIC FEE AMOUNT = |              |  |  | \$ 300.00<br><br><br><br><br><br><br><br><br><br><br>\$ 200.00<br><br><br><br><br><br><br><br><br><br><br>\$ 400.00<br><br><br><br><br><br><br><br><br><br><br>\$ 900.00 |              |
| Surcharge of \$130.00 for furnishing the oath or declaration later than 30 months from the earliest claimed priority date (37 CFR 1.492(e)).   |              |  |  |  |              |
| CLAIMS   | NUMBER FILED | NUMBER EXTRA                                       | RATE   |  |              |
| TOTAL  | 1 -20 =      | 0  | X \$ 50.00   |  |              |
| INDEPENDENT  | 1 - 3 =      | 0  | X \$ 200.00  |  |              |
| Multiple dependent claims(s) (if applicable)   |              |  | + \$360.00   |  |              |
| TOTAL OF ABOVE CALCULATIONS =  |              |  |  | \$ 900.00  |              |
| National Stage Application Size Fee-for each additional 50 sheets that exceeds 100 sheets \$ 250.00.   |              |  |  |  |              |
| Reduction by 1/2 for filing by small entity, if applicable. (Note 37 CFR 1.9, 1.27, 1.28).   |              |  |  | \$ -450.00   |              |
| SUBTOTAL =   |              |  |  | \$ 450.00  |              |
| Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 CFR 1.492(f)). +  |              |  |  |  |              |
| TOTAL NATIONAL FEE =   |              |  |  | \$ 450.00  |              |
| Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property +   |              |  |  | \$ 40.00   |              |
| TOTAL FEES ENCLOSED =  |              |  |  | \$ 490.00  |              |
|  |              |  |  | Amount to be:<br>refunded<br>charged   | \$<br>\$     |
| a. <input checked="" type="checkbox"/> A check in the amount of \$490.00 to cover the above fees is enclosed. SMALL ENTITY (\$150.00 for basic national stage fee; \$200.00 for national stage search fee; \$100.00 for national stage examination fee and \$40.00 for the assignment recordation fee).<br><br>b. <input type="checkbox"/> Please charge my Deposit Account No. 50-2866 in the amount of \$_____ to cover the above fees.<br><br>c. <input checked="" type="checkbox"/> The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 50-2866.   |              |  |  |  |              |
| NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed to request that the application be restored to pending status.   |              |  |  |  |              |
| Send All Correspondence To:  |              |  | WESTERMAN, HATTORI, DANIELS & ADRIAN, LLP<br>1250 Connecticut Avenue NW, Suite 700<br>Washington, DC 20036<br>CUSTOMER NO. 38834 |  |              |
| Typed or Printed Name: NEW Ken-Ichi Hattori  |              |  | Reg. No.: 32,861   |  |              |
| Signature   |              |  | Date: March 2, 2006  |  |              |

KH/yap